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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/616,251	07/08/2003	Peter Martin	484	9476
JOHN R. ROSS	7590 09/24/2008		EXAM	INER
TREX ENTERPRISES			YANG, NELSON C	
10455 PACIFIC SAN DIEGO, C	C CENTER CT. CA 92121		ART UNIT PAPER NUMBER	
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			09/24/2008	PAPER

Please find below and/or attached an Office communication concerning this application or proceeding.

The time period for reply, if any, is set in the attached communication.

Z. aal	Application No.	Applicant(s)	
Suppl.	10/616,251 MARTIN ET AL.		
Notice of Allowability	Examiner	Art Unit	
	Nelson Yang	1641 ′	
The MAILING DATE of this communication appears All claims being allowable, PROSECUTION ON THE MERITS IS herewith (or previously mailed), a Notice of Allowance (PTOL-85) NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RIOF the Office or upon petition by the applicant. See 37 CFR 1.313	(OR REMAINS) CLOSED in or other appropriate communing IGHTS. This application is subsequently and MPEP 1308.	this application. If not included nication will be mailed in due course.	
<u>_</u>			•
2. March The allowed claim(s) is/are <u>1-27,29-34, 36-47, renumbered</u>	<u>d 1-45</u> .		
3. Acknowledgment is made of a claim for foreign priority unall All b) Some* c) None of the: 1. Certified copies of the priority documents have 2. Certified copies of the priority documents have 3. Copies of the certified copies of the priority documents have International Bureau (PCT Rule 17.2(a)). * Certified copies not received: Applicant has THREE MONTHS FROM THE "MAILING DATE" noted below. Failure to timely comply will result in ABANDONM THIS THREE-MONTH PERIOD IS NOT EXTENDABLE. 4. A SUBSTITUTE OATH OR DECLARATION must be subminformal part application (PTO-152) which give 5. CORRECTED DRAWINGS (as "replacement sheets") must be comply including changes required by the Notice of Draftspers 1) hereto or 2) to Paper No./Mail Date	e been received. e been received in Application cuments have been received of this communication to file attached. Note the attached EXA es reason(s) why the oath or st be submitted.	No in this national stage application from a reply complying with the requirement MINER'S AMENDMENT or NOTICE of	nts
(b) ☐ including changes required by the attached Examiner's Paper No./Mail Date	s Amendment / Comment or	n the Office action of	
Identifying indicia such as the application number (see 37 CFR 1 each sheet. Replacement sheet(s) should be labeled as such in the			f
6. DEPOSIT OF and/or INFORMATION about the depo- attached Examiner's comment regarding REQUIREMENT			:
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Attachment(s) 1. ☐ Notice of References Cited (PTO-892)	5. ☐ Notice of Info	ormal Patent Application	
2. Notice of Draftperson's Patent Drawing Review (PTO-948)	6. Interview Su	nmary (PTO-413),	
3. ⊠ Information Disclosure Statements (PTO/SB/08),		fail Datemendment/Comment	
Paper No./Mail Date <u>4/21/08</u> 4. ☐ Examiner's Comment Regarding Requirement for Deposit	8. ⊠ Examiner's S	statement of Reasons for Allowance	
of Biological Material			

9. Other _____

DETAILED ACTION

Election/Restrictions

Claims 1-5, 7-17, 21-26, 29, 30, 38-47 are allowable. Claims 6, 18-20, and 27, previously withdrawn from consideration as a result of a restriction requirement, requires all the limitations of an allowable claim. Pursuant to the procedures set forth in MPEP § 821.04(a), the restriction requirement between the different species of light sources, as set forth in the Office action mailed on June 15, 2006, is hereby withdrawn and claims 6, 18-20, and 27 are hereby rejoined and fully examined for patentability under 37 CFR 1.104. In view of the withdrawal of the restriction requirement, applicant(s) are advised that if any claim presented in a continuation or divisional application is anticipated by, or includes all the limitations of, a claim that is allowable in the present application, such claim may be subject to provisional statutory and/or nonstatutory double patenting rejections over the claims of the instant application. Once the restriction requirement is withdrawn, the provisions of 35 U.S.C. 121 are no longer applicable. See *In re Ziegler*, 443 F.2d 1211, 1215, 170 USPQ 129, 131-32 (CCPA 1971). See also MPEP § 804.01.

Claims 1-5, 7-17, 21-26, 29, 30, 38-47 are directed to an allowable product. Pursuant to the procedures set forth in MPEP § 821.04(B), claims 31-37, directed to the process of making or using an allowable product, previously withdrawn from consideration as a result of a restriction requirement, are hereby rejoined and fully examined for patentability under 37 CFR 1.104.

Because all claims previously withdrawn from consideration under 37 CFR 1.142 have been rejoined, the restriction requirement as set forth in the Office action mailed on July 15, 2006 is hereby withdrawn. In view of the withdrawal of the restriction requirement as to the

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In claim 35, please change "The method as in claim 30" to "The method as in claim 34".

Please cancel claim 35, and amend claims 1, 31, 38, 41, and 42 as follows:

- 1. An optical sensor for monitoring molecular binding interactions, said sensor comprising:
- A) at least one porous silicon region comprising more than 1000 pores, each pore having a nominal width and a nominal depth at least 10 times larger than said nominal width, with the depth of each pore being approximately equal to the depth of at least most other pores in said porous silicon region, said porous silicon region defining a top surface and a bottom surface, and said bottom surface being parallel or approximately parallel to said top surface:
- B) at least one buffer-sample fluid flow channel located above said at least one porous silicon region providing a fluid flow passage across said porous silicon region;
- C) at least one light source for illuminating said at least one porous silicon region;
- D) at least one interference monitor adapted to monitor interference patterns caused by interference of light reflected from said top surface with light reflected from said bottom surface of said at least one porous silicon region, said interference monitor comprising a deep well linear photodiode array of pixels, each pixel having a photoelectron full well capacity of about 156 million photoelectrons or more, and having a frame rate of about one hundred or more frames of interference fringe data per second;
- E) a fluid flow control system for producing controlled flow of buffer solutions, ligand containing solutions, and analyte containing solutions through said at least one fluid flow channel; and
- F) a computer processor programmed with a computer program for making causing said processor to execute molecular binding measurements based on changes in the interference

patterns monitored by the at least one interference monitor while analytes bind with and disassociate from ligands attached to surfaces of said pores, said computer program comprising a special correlation method executable instructions for calculating optical path differences in measured interference fringe patterns wherein the measured fringe patterns is that are correlated to a test fringe pattern, wherein the test fringe pattern varies sinusoidally as a function of optical path differences divided by the wavelength of said light.

- 31. A method for measuring molecular binding interactions, utilizing an optical sensor comprising: having:
- a) at least one porous silicon region comprising more than 1000 pores, each pore having a nominal width and a nominal depth at least 10 times larger than said nominal width, with the depth of each pore being approximately equal to the depth of at least most other pores in said porous silicon region, said porous silicon region defining a top surface and a bottom surface, and said bottom surface being parallel or approximately parallel to said top surface;
- b) at least one buffer-sample fluid flow channel located above said at least one porous silicon region providing a fluid flow passage across said porous silicon region;
- c) at least one light source for illuminating said at least one porous silicon region;
- d) at least one spectral interference monitor for adapted to monitoring interference fringe patterns caused by interference of light reflected from said top surface with light reflected from and said bottom surface of said at least one porous silicon region, said interference monitor comprising a deep well linear photodiode array of pixels, each pixel having a photoelectron full well capacity

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of about 156 million photoelectrons or more, and having a frame rate of about one hundred or more frames of interference fringe data per second;

- e) a fluid flow control system for producing controlled flow of buffer solutions, ligand containing solutions, and analyte containing solutions through said at least one fluid flow channel; and
- f) a computer processor programmed with a computer program for making causing said

 processor to execute kinetic binding measurements based on changes in the spectral interference

 patterns monitored by the at least one interference monitor while analytes bind with and

 disassociate from ligands attached to surfaces of said pores, said computer program comprising

 :a special correlation method executable instructions for calculating optical path differences in

 measured interference fringe patterns monitored by said at least one spectral monitor while

 analytes bind with and disassociate from ligands attached to surfaces of said pores that are

 correlated the measured interference fringe patterns to a test fringe pattern, wherein the test

 fringe pattern varies sinusoidally as a function of optical path differences divided by the

 wavelength of said light, wherein said method comprises:
- A) immobilizing ligand molecules within said pores;
- B) causing a solution containing analyte molecules to flow across said porous silicon region to permit analyte molecules to diffuse close to and become bound at least temporarily by to said ligand molecules to form interference fringe patterns;
- C) illuminating at least a portion of said porous silicon region so as to produce reflections from said bottom surface and said top surface; and

- D) monitoring changes in spectral interference fringe patterns produced by light reflected from said top and bottom surfaces in order to obtain information concerning binding reactions between said ligand and said analyte.
- 38. An optical sensor for monitoring molecular binding interactions, said sensor comprising:

 A) at least one porous silicon region comprising more than 1000 pores, each pore having a nominal width and a nominal depth at least 10 times larger than said nominal width, with the depth of each pore being approximately equal to the depth of at least most other pores in said porous silicon region, said porous silicon region defining a top surface and a bottom surface, and said bottom surface being parallel or approximately parallel to said top surface;
- B) at least one buffer-sample fluid flow channel located above said at least one porous silicon region providing a fluid flow passage across said porous silicon region;
- C) at least one light source for illuminating said at least one porous silicon region;
- D) at least one interference monitor adapted to monitor interference patterns caused by interference of light reflected from said top surface with light reflected from said bottom surface of said at least one porous silicon region, said interference monitor comprising a deep well linear photodiode array of pixels, each pixel having a photoelectron full well capacity of about 156 million photoelectrons or more, and having a frame rate of about one hundred or more frames of interference fringe data per second;
- E) a fluid flow control system for producing controlled flow of buffer solutions, ligand containing solutions, and analyte containing solutions through said at least one fluid flow channel; and

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F) a processor means programmed with a computer program for making causing said processor means to execute kinetic molecular binding measurements based on changes in the interference patterns monitored by the at least one interference monitor while analytes bind with and disassociate from ligands attached to surfaces of said pores, said computer program comprising a special correlation method executable instructions for calculating optical path differences in measured interference fringe patterns calculation of optical path differences from measured interference fringe patterns wherein each measured fringe pattern is that are correlated to a test fringe pattern, wherein the test fringe pattern varies sinusoidally as a function of optical path differences divided by the wavelength of said light.

- 41. An optical sensor for monitoring molecular binding interactions, said sensor comprising:
- A) at least one porous silicon region, said porous silicon region defining a top surface and a bottom surface, <u>and</u> said bottom surface being parallel or approximately parallel to said top surface;
- B) at least one buffer-sample fluid flow channel located above said at least one porous silicon region providing a fluid flow passage across said porous silicon region;
- C) at least one light source for illuminating said at least one porous silicon region;
- D) at least one interference monitor adapted to monitor interference patterns caused by interference of light reflected from said top surface with light reflected from said bottom surface of said at least one porous silicon region, said interference monitor comprising a deep well linear photodiode array of pixels, each pixel having a photoelectron full well capacity of about 156

million photoelectrons or more, and having a frame rate of about one hundred or more frames of interference fringe data per second;

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- E) a fluid flow control system for producing controlled flow of buffer solutions, ligand containing solutions, and analyte containing solutions through said at least one fluid flow channel; and
- F) a computer processor programmed with a computer program for making causing said processor to execute molecular binding measurements based on changes in the interference patterns monitored by the at least one interference monitor while analytes bind with and disassociate from ligands attached to surfaces of said pores, said computer program comprising :a special correlation method executable instructions for calculating optical path differences in measured interference fringe patterns calculation of optical path differences from measured interference fringe patterns wherein each measured fringe pattern is that are correlated to a test fringe pattern, wherein the test fringe pattern varies sinusoidally as a function of optical path differences divided by the wavelength of said light.
- 42. An optical sensor for monitoring molecular binding interactions, said sensor comprising: A) at least one porous silicon region comprising more than 1,000 pores, each pore having a nominal width and a nominal depth at least 10 times larger than said nominal width, with the depth of each pore being approximately equal to the depth of at least most other pores in said porous silicon region, said porous silicon region defining a top surface and a bottom surface, and said bottom surface being parallel or approximately parallel to said top surface;

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- B) at least one buffer-sample fluid flow channel located above said at least one porous silicon region providing a fluid flow passage across said porous silicon region;
- C) at least one light source for illuminating said at least one porous silicon region;
- D) at least one interference monitor adapted to monitor interference patterns caused by interference of light reflected from said top surface with light reflected from said bottom surface of said at least one porous silicon region, said interference monitor comprising a deep well linear photodiode array of pixels, each pixel having a photoelectron full well capacity of about 156 million photoelectrons or more, and having a frame rate of about one hundred or more frames of interference fringe data per second;
- E) a fluid flow control system for producing controlled flow of buffer solutions, ligand containing solutions, and analyte containing solutions through said at least one fluid flow channel; and
- F) a computer processor programmed with a computer program for making causing said processor to execute molecular concentration measurements based on changes in the interference patterns monitored by the at least one interference monitor while analytes bind with and disassociate from ligands attached to surfaces of said pores, said computer program comprising a special correlation method executable instructions for calculating optical path differences in measured interference fringe patterns calculation of optical path differences from measured interference fringe patterns wherein each measured fringe pattern is that are correlated to a test fringe pattern, wherein the test fringe pattern varies sinusoidally as a function of optical path differences divided by the wavelength of said light.

The following is an examiner's statement of reasons for allowance: the prior art fails to teach a test fringe pattern that varies sinusoidally as a function of optical path differences divided by the wavelength of the light.

Any comments considered necessary by applicant must be submitted no later than the payment of the issue fee and, to avoid processing delays, should preferably accompany the issue fee. Such submissions should be clearly labeled "Comments on Statement of Reasons for Allowance."

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Nelson Yang whose telephone number is (571)272-0826. The examiner can normally be reached on 8:30-5:00.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Long V. Le can be reached on (571)272-0823. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

rejoined inventions, applicant(s) are advised that if any claim presented in a continuation or divisional application is anticipated by, or includes all the limitations of, a claim that is allowable in the present application, such claim may be subject to provisional statutory and/or nonstatutory double patenting rejections over the claims of the instant application. Once the restriction requirement is withdrawn, the provisions of 35 U.S.C. 121 are no longer applicable. See *In re Ziegler*, 443 F.2d 1211, 1215, 170 USPQ 129, 131-32 (CCPA 1971). See also MPEP § 804.01.

EXAMINER'S AMENDMENT

An examiner's amendment to the record appears below. Should the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the issue fee.

Authorization for this examiner's amendment was given in a telephone interview with John Ross on August 11 and September 15, 2008. Support for the amendments can be found on p.23-26 of the specification.

Please amend the Brief Description of the Drawings in the specification as follows:

On p. 4, line 21, please change FIGS. 6A&B to FIGS. 6A-C.

On p. 4, line 24, please change FIG. 9 to FIG. 9A-F.

On p. 5, line 6, please change FIG. 15 to FIGS. 15A and 15B

In claims 32, 36, 37 please change "The method as in claim 27" to "The method as in claim 31". In claims 33 and 34, please change "The method as in claim 28" to "The method as in claim 32".

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/Nelson Yang/ Patent Examiner, Art Unit 1641